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## SEQUENCE LISTING

<110> Tsukahara, Kappei Tsuchiya, Mamiko Jigami, Yoshifumi Nakayama, Kenichi Umemura, Mariko Okamoto, Michiyo <120> METHOD OF SCREENING FOR COMPOUNDS THAT INHIBIT THE ENZYMATIC ACTIVITY OF GWT1 GENE PRODUCT <130> 082368-004400US <140> 10/536,935 <141> <150> PCT/JP03/14909 <151> 2003-11-21 <150> JP 2002-339418 <151> 2002-11-22 <160> 18 <170> FastSEQ for Windows Version 4.0 <210> 1 <211> 1497 <212> DNA <213> Saccharomyces cerevisiae <220> <221> CDS <222> (1)..(1494) <400> 1 atg gca aca gta cat cag aag aat atg tcg act tta aaa cag aga aaa 48 Met Ala Thr Val His Gln Lys Asn Met Ser Thr Leu Lys Gln Arg Lys 1 15 gag gac ttt gtg aca ggg ctc aat ggc ggt tct ata aca gaa att aac 96 Glu Asp Phe Val Thr Gly Leu Asn Gly Gly Ser Ile Thr Glu Ile Asn 20 gca gtg aca tca att gct ttg gta act tac ata tca tgg aac tta ttg 144 Ala Val Thr Ser Ile Ala Leu Val Thr Tyr Ile Ser Trp Asn Leu Leu 35 aaa aat too aac ott atg oot oot ggo att too ago gtg caa tac ata 192 Lys Asn Ser Asn Leu Met Pro Pro Gly Ile Ser Ser Val Gln Tyr Ile 50 55 60 att gat ttt gca ttg aac tgg gtt gct ttg ctt cta tct att act att 240

Ile Asp Phe Ala Leu Asn Trp Val Ala Leu Leu Ser Ile Thr Ile

tat gct agt gaa cca tac ctt cta aac acg cta ata ctg tta cct tgt

Tyr Ala Ser Glu Pro Tyr Leu Leu Asn Thr Leu Ile Leu Leu Pro Cys

70

65

		gca Ala														336
		ata Ile 115														384
gaa Glu	aaa Lys 130	aag Lys	ccg Pro	tat Tyr	att Ile	act Thr 135	gcg Ala	tat Tyr	cgt Arg	ggt Gly	ggg Gly 140	atg Met	ctt Leu	att Ile	ctg Leu	432
act Thr 145	gct Ala	att Ile	gcc Ala	atc Ile	ttg Leu 150	gct Ala	gta Val	gat Asp	ttt Phe	cca Pro 155	att Ile	ttc Phe	cca Pro	agg Arg	agg Arg 160	480
		aag Lys														528
gga Gly	tca Ser	ttc Phe	gtt Val 180	ttc Phe	agt Ser	aac Asn	ggt Gly	att Ile 185	gtt Val	tct Ser	tct Ser	agg Arg	gca Ala 190	ctg Leu	ttg Leu	576
		cta Leu 195														624
		tta Leu														672
ttg Leu 225	ttt Phe	ttt Phe	gta Val	aaa Lys	aat Asn 230	ttg Leu	gaa Glu	tat Tyr	caa Gln	gaa Glu 235	cat His	gtc Val	aca Thr	gaa Glu	tat Tyr 240	720
		cat His														768
ttg Leu	acc Thr	ttt Phe	att Ile 260	gat Asp	ccc Pro	gtc Val	aca Thr	aga Arg 265	atg Met	gtt Val	cca Pro	cgc Arg	tgc Cys 270	tca Ser	att Ile	816
gca Ala	ata Ile	ttc Phe 275	att Ile	tca Ser	tgc Cys	att Ile	tat Tyr 280	gaa Glu	tgg Trp	cta Leu	ctt Leu	tta Leu 285	aag Lys	gac Asp	gat Asp	864
cgc Arg	act Thr 290	tta Leu	aac Asn	ttt Phe	tta Leu	att Ile 295	ttg Leu	gct Ala	gat Asp	aga Arg	aat Asn 300	tgt Cys	ttc Phe	ttc Phe	agt Ser	912
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ctt Leu	tgg Trp	ggc Gly	caa Gln	aac Asn 325	acg Thr	gga Gly	ttt Phe	tac Tyr	ttg Leu 330	ttg Leu	gga Gly	aat Asn	aaa Lys	cca Pro 335	act Thr	1008

						cct Pro										1056
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tgc Cys	ttg Leu	act Thr	gac Asp 420	aaa Lys	att Ile	ttc Phe	ggt Gly	aat Asn 425	tct Ser	tcg Ser	gaa Glu	tat Tyr	tat Tyr 430	aaa Lys	gtt Val	1296
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						ggt Gly 455										1392
						tca Ser										1440
ttc Phe	ata Ile	gct Ala	gtc Val	ata Ile 485	tcg Ser	gtt Val	ttc Phe	ttg Leu	tat Tyr 490	aga Arg	aaa Lys	aga Arg	ata Ile	ttc Phe 495	att Ile	1488
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Lys Asn Ser Asn Leu Met Pro Pro Gly Ile Ser Ser Val Gln Tyr Ile Ile Asp Phe Ala Leu Asn Trp Val Ala Leu Leu Leu Ser Ile Thr Ile 75 Tyr Ala Ser Glu Pro Tyr Leu Leu Asn Thr Leu Ile Leu Leu Pro Cys 90 Leu Leu Ala Phe Ile Tyr Gly Lys Phe Thr Ser Ser Ser Lys Pro Ser 105 Asn Pro Ile Tyr Asn Lys Lys Lys Met Ile Thr Gln Arg Phe Gln Leu Glu Lys Lys Pro Tyr Ile Thr Ala Tyr Arg Gly Gly Met Leu Ile Leu 135 Thr Ala Ile Ala Ile Leu Ala Val Asp Phe Pro Ile Phe Pro Arg Arg 155 Phe Ala Lys Val Glu Thr Trp Gly Thr Ser Leu Met Asp Leu Gly Val 170 Gly Ser Phe Val Phe Ser Asn Gly Ile Val Ser Ser Arg Ala Leu Leu 185 Lys Asn Leu Ser Leu Lys Ser Lys Pro Ser Phe Leu Lys Asn Ala Phe 200 Asn Ala Leu Lys Ser Gly Gly Thr Leu Leu Phe Leu Gly Leu Leu Arg 215 Leu Phe Phe Val Lys Asn Leu Glu Tyr Gln Glu His Val Thr Glu Tyr 230 235 Gly Val His Trp Asn Phe Phe Ile Thr Leu Ser Leu Leu Pro Leu Val 245 250 Leu Thr Phe Ile Asp Pro Val Thr Arg Met Val Pro Arg Cys Ser Ile 260 265 270 Ala Ile Phe Ile Ser Cys Ile Tyr Glu Trp Leu Leu Lys Asp Asp Arg Thr Leu Asn Phe Leu Ile Leu Ala Asp Arg Asn Cys Phe Phe Ser Ala Asn Arg Glu Gly Ile Phe Ser Phe Leu Gly Tyr Cys Ser Ile Phe Leu Trp Gly Gln Asn Thr Gly Phe Tyr Leu Leu Gly Asn Lys Pro Thr 330 Leu Asn Asn Leu Tyr Lys Pro Ser Thr Gln Asp Val Val Ala Ala Ser 345 Lys Lys Ser Ser Thr Trp Asp Tyr Trp Thr Ser Val Thr Pro Leu Ser 360

Gly Leu Cys Ile Trp Ser Thr Ile Phe Leu Val Ile Ser Gln Leu Val 375 Phe Gln Tyr His Pro Tyr Ser Val Ser Arg Arg Phe Ala Asn Leu Pro 390 395 Tyr Thr Leu Trp Val Ile Thr Tyr Asn Leu Leu Phe Leu Thr Gly Tyr 405 Cys Leu Thr Asp Lys Ile Phe Gly Asn Ser Ser Glu Tyr Tyr Lys Val 425 Ala Glu Cys Leu Glu Ser Ile Asn Ser Asn Gly Leu Phe Leu Phe Leu 440 Leu Ala Asn Val Ser Thr Gly Leu Val Asn Met Ser Met Val Thr Ile 455 Asp Ser Ser Pro Leu Lys Ser Phe Leu Val Leu Leu Ala Tyr Cys Ser 470 475 Phe Ile Ala Val Ile Ser Val Phe Leu Tyr Arg Lys Arg Ile Phe Ile 485 490 Lys Leu <210> 3 <211> 1458 <212> DNA <213> Candida albicans <220> <221> CDS <222> (1)..(1455) <400> 3 atg tca tcg tct tta aaa caa ttg aaa gaa caa ttt gtc tca gat ttg 48 Met Ser Ser Leu Lys Gln Leu Lys Glu Gln Phe Val Ser Asp Leu act ggt ggc aca att gaa gaa att tat gct gta acc agt ata gca tta 96 Thr Gly Gly Thr Ile Glu Glu Ile Tyr Ala Val Thr Ser Ile Ala Leu tca tct tat ttg tcc ttt aga ttg ttg aaa aag tct ctt ggt gat tta 144 Ser Ser Tyr Leu Ser Phe Arg Leu Leu Lys Lys Ser Leu Gly Asp Leu gct ttg att tac gac tac att ctt aat gtg ttg aca att cta gca tcc 192 Ala Leu Ile Tyr Asp Tyr Ile Leu Asn Val Leu Thr Ile Leu Ala Ser 55 att act gtt tat agc aac agc cct tct tat ttg cat tat ttt att gtt 240 Ile Thr Val Tyr Ser Asn Ser Pro Ser Tyr Leu His Tyr Phe Ile Val att cca tca tta gtt ata tat cta gtg aat tac cat gtt gag aaa cca 288 Ile Pro Ser Leu Val Ile Tyr Leu Val Asn Tyr His Val Glu Lys Pro 85 90

														gac Asp		336
														atg Met		384
														ttc Phe		432
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														agt Ser		576
														ctt Leu		624
tta Leu	gga Gly 210	gct Ala	att Ile	cgt Arg	ttt Phe	gtt Val 215	agt Ser	gtt Val	aag Lys	caa Gln	ttg Leu 220	gac Asp	tat Tyr	cag Gln	gaa Glu	672
cac His 225	gaa Glu	aca Thr	gag Glu	tat Tyr	gga Gly 230	atc Ile	cat His	tgg Trp	aat Asn	ttt Phe 235	ttc Phe	ttc Phe	aca Thr	tta Leu	ggg Gly 240	720
ttc Phe	ttg Leu	cca Pro	att Ile	gta Val 245	ttg Leu	gga Gly	ata Ile	tta Leu	gac Asp 250	ccg Pro	gtg Val	ttg Leu	aat Asn	ttg Leu 255	gtt Val	768
cca Pro	cgc Arg	ttc Phe	ata Ile 260	ata Ile	gga Gly	att Ile	ggt Gly	atc Ile 265	tca Ser	att Ile	gct Ala	tat Tyr	gag Glu 270	gta Val	gcg Ala	816
ttg Leu	aat Asn	aag Lys 275	act Thr	ggt Gly	ttg Leu	ttg Leu	aag Lys 280	ttc Phe	att Ile	ttg Leu	agc Ser	agc Ser 285	gaa Glu	aac Asn	aga Arg	864
														ttt Phe		912
gga Gly 305	tat Tyr	ctt Leu	tgt Cys	att Ile	ttt Phe 310	ata Ile	att Ile	ggt Gly	cag Gln	tct Ser 315	ttt Phe	ggg Gly	tca Ser	ttt Phe	gtt Val 320	960
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cgt att tca aaa a Arg Ile Ser Lys I 340					
gtc gcc act act c Val Ala Thr Thr G 355	ln Gly Leu				
gct ttc agt ttg t Ala Phe Ser Leu P 370					
aga cga ttg gcc a Arg Arg Leu Ala A 385				Ser Tyr A	
gct acg ttt tta t Ala Thr Phe Leu L 4			Glu Lys Phe		
aac ctt act tct a Asn Leu Thr Ser T 420					
atc ttc ttg gtc a Ile Phe Leu Val S 435	er Asn Leu				
aac act ttg gaa a Asn Thr Leu Glu T 450					
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Ala Leu Ile Tyr A 50	sp Tyr Ile : 55	Leu Asn Val	Leu Thr Ile 60	Leu Ala Se	er
Ile Thr Val Tyr S	er Asn Ser	Pro Ser Tyr	Leu His Tyr	Phe Ile Va	al

Ile Pro Ser Leu Val Ile Tyr Leu Val Asn Tyr His Val Glu Lys Pro 85 90 Ser Ser Pro His Arg Gln Asn Asp Thr Lys Glu Asp Lys Ser Asp Glu 105 Leu Leu Pro Arg Lys Gln Phe Ile Thr Ala Tyr Arg Ser Gln Met Leu Ile Ile Thr Asn Leu Ala Ile Leu Ala Val Asp Phe Pro Ile Phe Pro 135 140 Arg Arg Phe Ala Lys Val Glu Thr Trp Gly Thr Ser Met Met Asp Leu 150 155 Gly Val Gly Ser Phe Val Phe Ser Met Gly Leu Ala Asn Ser Arg Gln 165 170 Leu Ile Lys Asn His Thr Asp Asn Tyr Lys Phe Ser Trp Lys Ser Tyr 185 Leu Lys Thr Ile Lys Gln Asn Phe Ile Lys Ser Val Pro Ile Leu Val Leu Gly Ala Ile Arg Phe Val Ser Val Lys Gln Leu Asp Tyr Gln Glu His Glu Thr Glu Tyr Gly Ile His Trp Asn Phe Phe Thr Leu Gly 235 Phe Leu Pro Ile Val Leu Gly Ile Leu Asp Pro Val Leu Asn Leu Val 250 Pro Arg Phe Ile Ile Gly Ile Gly Ile Ser Ile Ala Tyr Glu Val Ala 265 Leu Asn Lys Thr Gly Leu Leu Lys Phe Ile Leu Ser Ser Glu Asn Arg Leu Glu Ser Leu Ile Thr Met Asn Lys Glu Gly Ile Phe Ser Phe Ile 295 Gly Tyr Leu Cys Ile Phe Ile Ile Gly Gln Ser Phe Gly Ser Phe Val 310 315 Leu Thr Gly Tyr Lys Thr Lys Asn Asn Leu Ile Thr Ile Ser Lys Ile 325 Arg Ile Ser Lys Lys Gln His Lys Lys Glu Leu Leu Phe Phe Ser 340 Val Ala Thr Thr Gln Gly Leu Tyr Leu Ala Cys Ile Phe Tyr His Leu Ala Phe Ser Leu Phe Ile Ser Asn Leu Ser Phe Leu Gln Pro Ile Ser 375 Arg Arg Leu Ala Asn Phe Pro Tyr Val Met Trp Val Val Ser Tyr Asn

385					390					395					400	
Ala	Thr	Phe	Leu	Leu 405	Cys	Tyr	Asp	Leu	Ile 410	Glu	Lys	Phe	Ile	Pro 415	Gly	
Asn	Leu	Thr	Ser 420	Thr	Val	Leu	Asp	Ser 425		Asn	Asn	Asn	Gly 430	Leu	Phe	
Ile	Phe	Leu 435	Val	Ser	Asn	Leu	Leu 440	Thr	Gly	Phe	Ile	Asn 445	Met	Ser	Ile	
Asn	Thr 450	Leu	Glu	Thr	Ser	Asn 455	Lys	Met	Ala	Val	Ile 460	Ile	Leu	Ile	Gly	
Tyr 465	Ser	Leu	Thr	Trp	Thr 470	Leu	Leu	Ala	Leu	Tyr 475	Leu	Asp	Lys	Arg	Lys 480	
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			aca Thr 20													96
			ttg Leu												tta Leu	_ 144
			tac Tyr													192
			tat Tyr													240
att Ile	cca Pro	tca Ser	tta Leu	gtt Val 85	ata Ile	tat Tyr	cta Leu	gtg Val	aat Asn 90	tac Tyr	cat His	gtt Val	gag Glu	aaa Lys 95	cca Pro	288
			cat His 100													336
			aga Arg													384

		115					120					125				
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aga Arg 145	aga Arg	ttt Phe	gcc Ala	aaa Lys	gtg Val 150	gaa Glu	aca Thr	tgg Trp	ggc Gly	acg Thr 155	tca Ser	atg Met	atg Met	gat Asp	tta Leu 160	480
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cca Pro	cgc Arg	ttc Phe	ata Ile 260	ata Ile	gga Gly	att Ile	ggt Gly	atc Ile 265	tca Ser	att Ile	ggt Gly	tat Tyr	gag Glu 270	gta Val	gcg Ala	816
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						aag Lys										1008
cgt Arg	att Ile	tca Ser	aaa Lys 340	aaa Lys	caa Gln	cac His	aag Lys	aaa Lys 345	gag Glu	ctg Leu	ctg Leu	ctg Leu	ttt Phe 350	ttc Phe	tca Ser	1056
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aac ctt act Asn Leu Thr								1296
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tat agt ctt Tyr Ser Leu 465	act tgg ac Thr Trp Th 47	r Leu Leu	gcc tta Ala Leu	tat ttg Tyr Leu 475	gat aag Asp Lys	agg Arg	aag Lys 480	1440
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Leu Leu Pro Arg Lys Gln Phe Ile Thr Ala Tyr Arg Ser Gln Met Leu 120 Ile Ile Thr Asn Leu Ala Ile Leu Ala Val Asp Phe Pro Ile Phe Pro 135 Arg Arg Phe Ala Lys Val Glu Thr Trp Gly Thr Ser Met Met Asp Leu 150 155 Gly Val Gly Ser Phe Val Phe Ser Met Gly Leu Ala Asn Ser Arg Gln 170 Leu Ile Lys Asn His Thr Asp Asn Tyr Lys Phe Ser Trp Lys Ser Tyr Leu Lys Thr Ile Lys Gln Asn Phe Ile Lys Ser Val Pro Ile Leu Val 200 Leu Gly Ala Ile Arg Phe Val Ser Val Lys Gln Leu Asp Tyr Gln Glu 215 His Glu Thr Glu Tyr Gly Ile His Trp Asn Phe Phe Thr Leu Gly 230 235 Phe Leu Pro Ile Val Leu Gly Ile Leu Asp Pro Val Leu Asn Leu Val 245 Pro Arg Phe Ile Ile Gly Ile Gly Ile Ser Ile Gly Tyr Glu Val Ala 265 Leu Asn Lys Thr Gly Leu Leu Lys Phe Ile Leu Ser Ser Glu Asn Arg Leu Glu Ser Leu Ile Ala Met Asn Lys Glu Gly Ile Phe Ser Phe Ile 295 Gly Tyr Leu Cys Ile Phe Ile Ile Gly Gln Ser Phe Gly Ser Phe Val 315 320 Leu Thr Gly Tyr Lys Thr Lys Asn Asn Leu Ile Thr Ile Ser Lys Ile 330 Arg Ile Ser Lys Lys Gln His Lys Lys Glu Leu Leu Phe Phe Ser 345 Val Ala Thr Thr Gln Gly Leu Tyr Leu Ala Cys Ile Phe Tyr His Leu Ala Phe Ser Leu Phe Ile Ser Asn Leu Ser Phe Leu Gln Pro Ile Ser 375 380 Arg Arg Leu Ala Asn Phe Pro Tyr Val Met Trp Val Val Ser Tyr Asn Ala Thr Phe Leu Cys Tyr Asp Leu Ile Glu Lys Phe Ile Pro Gly 405 Asn Leu Thr Ser Thr Val Leu Asp Ser Ile Asn Asn Asn Gly Leu Phe 420 425 Ile Phe Leu Val Ser Asn Leu Leu Thr Gly Phe Ile Asn Met Ser Ile

435		440	445	
Asn Thr Leu Glu 450	Thr Ser Asn 455	Lys Met Ala V	Val Ile Ile Leu : 460	Ile Gly
Tyr Ser Leu Thr 465	Trp Thr Leu 470		Tyr Leu Asp Lys <i>i</i> 475	Arg Lys 480
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aac gtt ttg tgg Asn Val Leu Trp 35	gta aac atg Val Asn Met	act gcg aga a Thr Ala Arg A 40	aac atc tta ccc a Asn Ile Leu Pro I 45	aaa ggg 144 Lys Gly
			ett tgc tta att o Phe Cys Leu Ile F 60	
Phe Val Ile Tyr	gtt tca tcg Val Ser Ser 70	aaa gtt ggc o Lys Val Gly V	gtt ttc act ctt t /al Phe Thr Leu ( 75	cgc ata 240 Cys Ile 80
gcc tct ttt ttg Ala Ser Phe Leu	cct tcc ttc Pro Ser Phe 85	gtc ctt cat o Val Leu His V 90	gtt ata agt cca a /al Ile Ser Pro l	att aat 288 [le Asn 95
tgg gat gtg ctg Trp Asp Val Leu 100	aga aga aaa Arg Arg Lys	cct ggt tgt t Pro Gly Cys C 105	gt ctt act aaa a Cys Leu Thr Lys I 110	aaa aat 336 Lys Asn
gaa aat act ttt Glu Asn Thr Phe 115	gat cga cga Asp Arg Arg	att gct gga g Ile Ala Gly V 120	gtc aca ttt tat c Val Thr Phe Tyr A 125	egt tct 384 Arg Ser
caa atg atg ttg Gln Met Met Leu 130	gtt act gtc Val Thr Val 135	act tgc atc o Thr Cys Ile I	etg gcc gtt gac t Leu Ala Val Asp E 140	ett acc 432 Phe Thr
ctt ttc ccg agg Leu Phe Pro Arg	aga tat gcc Arg Tyr Ala	aaa gtt gaa a Lys Val Glu T	acc tgg gga aca t hr Trp Gly Thr S	cca ctg 480 Ser Leu

145					150					155					160	
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					att Ile											576
tgg Trp	aat Asn	tct Ser 195	ttc Phe	atc Ile	ctt Leu	ttg Leu	att Ile 200	tta Leu	gga Gly	ttt Phe	gcg Ala	cgc Arg 205	atg Met	ttt Phe	tta Leu	624
					tac Tyr											672
					acc Thr 230											720
					aaa Lys											768
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ggt Gly	att Ile 290	gct Ala	tct Ser	ctt Leu	ccc Pro	gga Gly 295	tac Tyr	att Ile	gct Ala	att Ile	tac Tyr 300	ttt Phe	tat Tyr	gga Gly	atg Met	912
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cgc Arg	cga Arg	ctt Leu 355	gct Ala	aat Asn	acg Thr	cct Pro	tat Tyr 360	gtt Val	gcg Ala	aat Asn	gtt Val	gcc Ala 365	ttt Phe	atc Ile	aat Asn	1104
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					gga Gly 390											1200

aat aat Asn Asn															1248
gta gtt Val Val															1296
ttg aca Leu Thr															1344
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Asn Leu 50	Gly	Phe	Leu	Val	Glu 55	Phe	Phe	Ile	Phe	Cys 60	Leu	Ile	Pro	Leu	
Phe Val 65	Ile	Tyr	Val	Ser 70	Ser	Lys	Val	Gly	Val 75	Phe	Thr	Leu	Cys	Ile 80	
Ala Ser	Phe	Leu	Pro 85	Ser	Phe	Val	Leu	His 90	Val	Ile	Ser	Pro	Ile 95	Asn	
Trp Asp	Val	Leu 100	Arg	Arg	Lys	Pro	Gly 105	Cys	Cys	Leu	Thr	Lys 110	Lys	Asn	
Glu Asn	Thr 115	Phe	Asp	Arg	Arg	Ile 120	Ala	Gly	Val	Thr	Phe 125	Tyr	Arg	Ser	
Gln Met 130	Met	Leu	Val	Thr	Val 135	Thr	Cys	Ile	Leu	Ala 140	Val	Asp	Phe	Thr	
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Gly Arg	Lys	Asn 180	Asp	Ile	Lys	Lys	Pro 185	Asn	Ala	Phe	Lys	Asn 190	Val	Leu	
Trp Asn	Ser	Phe	Ile	Leu	Leu	Ile	Leu	Gly	Phe	Ala	Arg	Met	Phe	Leu	

Thr Lys Ser Ile Asn Tyr Gln Glu His Val Ser Glu Tyr Gly Met His 215 Trp Asn Phe Phe Thr Leu Gly Phe Met Ala Leu Gly Val Phe Phe 230 235 Phe Arg Arg Ser Leu Lys Lys Val Ser Tyr Phe Asn Leu Ala Thr Phe 245 250 Ile Thr Leu Leu His His Cys Leu Leu Val Leu Thr Pro Phe Gln Lys 265 Trp Ala Leu Ser Ala Pro Arg Thr Asn Ile Leu Ala Gln Asn Arg Glu 275 Gly Ile Ala Ser Leu Pro Gly Tyr Ile Ala Ile Tyr Phe Tyr Gly Met Tyr Thr Gly Ser Val Val Leu Ala Asp Arg Pro Leu Met Tyr Thr Arg 315 Ala Glu Ser Trp Lys Arg Phe Gln Arg Leu Leu Phe Pro Leu Cys Ile 330 Leu Leu Val Leu Tyr Leu Val Ser Asn Phe Leu Ser Val Gly Val Ser 345 Arg Arg Leu Ala Asn Thr Pro Tyr Val Ala Asn Val Ala Phe Ile Asn 360 Met Phe Phe Leu Thr Ile Tyr Ile Leu Ile Asp Ala Tyr Leu Phe Pro Ser Ser Val Pro Tyr Gly Ser Arg Val Pro Lys Leu Leu Glu Asp Ala 390 Asn Asn Gly Leu Leu Val Phe Leu Ile Ala Asn Val Leu Thr Gly

405 410 415

Val Val Asn Leu Ser Phe Asp Thr Leu His Ser Ser Asn Ala Lys Gly 420 425 430

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<220>

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aac Asn 25	gcc Ala	gtc Val	acc Thr	ttg Leu	gtt Val 30	gct Ala	tcg Ser	gta Val	tcc Ser	gtt Val 35	ttt Phe	ctg Leu	tgg Trp	tca Ser	att Ile 40	150
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aca Thr	tat Tyr	cgc Arg	gcc Ala 140	gcc Ala	atg Met	atg Met	gtt Val	atc Ile 145	acg Thr	tgc Cys	atc Ile	gct Ala	atc Ile 150	ttg Leu	gct Ala	486
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	gtc Val															966
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	cag Gln															1110
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	tgg Trp				Phe	Asn		Ala	Gln	Leu		Val				1254
	gaa Glu 410															1302
agc Ser 425	gaa Glu	tct Ser	gag Glu	cga Arg	gtc Val 430	gat Asp	ttt Phe	gct Ala	acg Thr	agc Ser 435	cga Arg	atc Ile	atg Met	tcg Ser	gcc Ala 440	1350
	aat Asn															1398
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gcc Ala	atc Ile	gct Ala	gtt Val	ctc Leu	att Ile	gga Gly	tat Tyr	tca Ser	tcc Ser	att Ile	atc Ile	aca Thr	ggg Gly	gtt Val	gct Ala	1494

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400

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476

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Thr Pro Tyr Ser Ala Ala Ala Leu Leu Val Asp Phe Leu Leu Asn Val 5.5

Leu Ala Ile Leu Phe Ala Thr Thr Leu Tyr Ser Ser Ala Pro Leu Leu 65 70 75 80

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Lys Arg Pro Arg Thr Pro Val Lys Ala Lys Pro Pro Arg Gln Ser Ala 110

Arg Ala Gly Lys Asp Asp Ser Lys His Ala Thr Ala Leu Pro Glu Ser

Leu Pro Ile His Pro Phe Leu Thr Thr Tyr Arg Ala Ala Met Met Val

Ile Thr Cys Ile Ala Ile Leu Ala Val Asp Phe Arg Ile Phe Pro Arg 155

Arg Phe Ala Lys Val Glu Asn Trp Gly Thr Ser Leu Met Asp Leu Gly 165 170

Val Gly Ser Phe Val Phe Ser Gly Gly Val Val Ser Ala Arg Ser Leu 185

Leu Lys Ser Arg Thr Asn Gly Ser Lys Arg Leu Pro Leu Ala Lys Arg 200 205

Leu Ile Ala Ser Thr Arg His Ser Ile Pro Leu Leu Val Leu Gly Leu 215

Ile Arg Leu Tyr Ser Val Lys Gly Leu Asp Tyr Ala Glu His Val Thr 230 235

Glu Tyr Gly Val His Trp Asn Phe Phe Phe Thr Leu Gly Leu Leu Pro 245 250 Pro Phe Val Glu Val Phe Asp Ala Leu Ala Thr Ile Ile Pro Ser Tyr 265 Glu Val Leu Ser Val Gly Ile Ala Val Leu Tyr Gln Val Ala Leu Glu 280 Ser Thr Asp Leu Lys Ser Tyr Ile Leu Val Ser Pro Arg Gly Pro Ser 295 Leu Leu Ser Lys Asn Arg Glu Gly Val Phe Ser Phe Ser Gly Tyr Leu Ala Ile Phe Leu Ala Gly Arg Ala Ile Gly Ile Arg Ile Ile Pro Arg 330 Gly Thr Ser Phe Ser Arg Ser Pro Glu Gln Ala Arg Arg Arg Val Leu 345 Ile Ser Leu Gly Val Gln Ala Leu Val Trp Thr Thr Leu Phe Val Leu 360 Asn Ser Thr Tyr Ala Met Gly Tyr Gly Ala Asn Ile Pro Val Ser Arg Arg Leu Ala Asn Met Pro Tyr Val Leu Trp Val Ser Ala Phe Asn Thr 390 395 Ala Gln Leu Phe Val Phe Cys Leu Ile Glu Thr Leu Cys Phe Pro Ala 405 410 Val His Arg Thr Thr Gln Glu Ser Glu Ser Glu Arg Val Asp Phe 425 Ala Thr Ser Arg Ile Met Ser Ala Phe Asn Lys Asn Ser Leu Ala Ile 435 Phe Leu Leu Ala Asn Leu Leu Thr Gly Ala Val Asn Leu Ser Ile Ser 455 Thr Ile Asp Ala Asn Thr Ala Gln Ala Ile Ala Val Leu Ile Gly Tyr Ser Ser Ile Ile Thr Gly Val Ala Leu Ala Leu His His Ala Asn Ile 490 Lys Val Leu Pro Phe 500 <210> 11

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						agg Arg										735
						cct Pro										783
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cgc t Arg T																1317
tta a Leu T																1365
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tcc a Ser I 420	ata [le	ttt Phe	ttg Leu	atc Ile	ggc Gly 425	ttg Leu	tct Ser	att Ile	gga Gly	gat Asp 430	cat His	gtt Val	tta Leu	agg Arg	ctc Leu 435	1461
agt t Ser L	ta Leu	cca Pro	cca Pro	aga Arg 440	aga Arg	gag Glu	agg Arg	gtc Val	gtg Val 445	tca Ser	gaa Glu	aca Thr	aat Asn	gaa Glu 450	gag Glu	1509
cat g His G	gag Slu	cag Gln	agt Ser 455	cat His	ttt Phe	gag Glu	aga Arg	aaa Lys 460	aaa Lys	ttg Leu	gat Asp	ttg Leu	att Ile 465	atg Met	gag Glu	1557
ttg a Leu I	le	gga Gly 470	tat Tyr	agc Ser	tta Leu	ggc Gly	tgg Trp 475	tgg Trp	gca Ala	ctc Leu	tta Leu	gga Gly 480	ggc Gly	tgg Trp	att Ile	1605
tgg g Trp A										gtaa	agtgo	gac a	atctt	tggt	a	1655
atatt	gta	cc t	atac	ctaat	c cc	etgca	taaa	gg	icc e	ac c	gct d	ct t	at o	gta t	tt	1707

## Ala Asn Ala Pro Tyr Val Phe 495 500

Trp Val Ala Ala Tyr Asn Thr Thr Phe Leu Leu Gly Tyr Leu Leu 505 510 515	1755
acc cac att att cca tct ccc acc tct tcc caa aca tca cca tcg atc Thr His Ile Ile Pro Ser Pro Thr Ser Ser Gln Thr Ser Pro Ser Ile 520 525 530	1803
tta gtg cct ccc ttg ctc gac gct atg aat aaa aac ggt ctc gcg ata Leu Val Pro Pro Leu Leu Asp Ala Met Asn Lys Asn Gly Leu Ala Ile 535 540 545	1851
ttt ttg gcg gcc aac ttg ctt aca gga ctg gtg aat gtg agc atg aag Phe Leu Ala Ala Asn Leu Leu Thr Gly Leu Val Asn Val Ser Met Lys 550 555 560	1899
aca atg tat gcg ccg gcg tgg ttg tca atg ggg gtg tta atg ttg tat Thr Met Tyr Ala Pro Ala Trp Leu Ser Met Gly Val Leu Met Leu Tyr 565 570 580	1947
acc ttg aca atc agt tgt gta ggg tgg ata ctg aaa gga cgg agg atc Thr Leu Thr Ile Ser Cys Val Gly Trp Ile Leu Lys Gly Arg Arg Ile 585 590 595	1995
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												tca Ser				336
												gca Ala 125				384
												tcg Ser				432
tta Leu 145	tct Ser	ccc Pro	gat Asp	ccg Pro	aca Thr 150	aca Thr	tcc Ser	ccc Pro	atg Met	tcg Ser 155	cca Pro	agt Ser	agt Ser	tct Ser	tca Ser 160	480
gct Ala	tca Ser	gga Gly	cat His	gaa Glu 165	gac Asp	cct Pro	ttg Leu	Gly ggg	att Ile 170	atg Met	ggc Gly	gtt Val	aac Asn	aga Arg 175	cgg Arg	528
agg Arg	tcg Ser	cta Leu	tta Leu 180	gaa Glu	gga Gly	gtt Val	tcg Ser	ctt Leu 185	gat Asp	gtt Val	ccg Pro	tca Ser	cat His 190	atc Ile	gac Asp	576
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Ser	Lys	Val 195	Arg	Ile	Ser	Pro	Val 200	Pro	Tyr	Leu	Arg	Leu 205	Lys	Lys	Ser	
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ttg Leu	gcg Ala	gta Val	gat Asp	ttt Phe 245	gaa Glu	gtg Val	ttt Phe	cct Pro	aga Arg 250	tgg Trp	cag Gln	ggc Gly	aag Lys	tgc Cys 255	gaa Glu	768
gat Asp	ttt Phe	ggt Gly	act Thr 260	agt Ser	ctg Leu	atg Met	gac Asp	gtg Val 265	ggt Gly	gtc Val	Gly ggg	tca Ser	ttc Phe 270	gtc Val	ttt Phe	816
tcc Ser	ctc Leu	ggt Gly 275	ctc Leu	gtc Val	tcc Ser	aca Thr	aaa Lys 280	tct Ser	ctt Leu	tct Ser	cct Pro	cca Pro 285	cct Pro	cca Pro	act Thr	864
cct Pro	acg Thr 290	ccc Pro	tcc Ser	tcg Ser	ccc Pro	gct Ala 295	ctc Leu	aac Asn	tct Ser	cac His	atc Ile 300	att Ile	ccc Pro	ctc Leu	acc Thr	912
ccg	tcc	ccg	ttc	act	tcc	atc	ctc	atc	tcg	ctc	cga	aaa	tcc	atc	ccc	960

Pro 305	Ser	Pro	Phe	Thr	Ser 310	Ile	Leu	Ile	Ser	Leu 315	Arg	Lys	Ser	Ile	Pro 320	
							cgg Arg									1008
							tac Tyr									1056
							ctc Leu 360									1104
							ctt Leu									1152
							ctc Leu									1200
						_	aac Asn	_	_				Ser			1248
							atc Ile									1296
							aga Arg 440									1344
							cat His									1392
att Ile 465	atg Met	gag Glu	Leu	Ile	Gly	Tyr	agc Ser	Leu	Gly	Trp	Trp	gca Ala	ctc Leu	Leu	gga Gly 480	1440
ggc Gly	tgg Trp	att Ile	tgg Trp	gcc Ala 485	ggc Gly	G]À aaa	gag Glu	gta Val	tcc Ser 490	agg Arg	cgt Arg	tta Leu	gcc Ala	aac Asn 495	gct Ala	1488
							gca Ala									1536
							att Ile 520									1584
							ccc Pro	_		_	_	_				1632
							gcc Ala									1680

545	550	555	560
	atg tat gcg ccg gcg Met Tyr Ala Pro Ala 570		
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Thr Tyr Ala Leu Trp	Ile Ala Leu Ser Pro	Tyr Ile Arg His Gly	Leu
35	40	45	
Leu Asn Asn Tyr Leu 50	Ile Cys Val Leu Pro 55	Leu Leu Phe Gly Val	Thr
Ile Phe Ser Thr Ser	Pro Leu Val Phe Thr 70	Ser Phe Leu Ser Ile	Ile
65		75	80
Ser Leu Ala Phe Ile	Thr Lys Ser Gln Lys	Cys Phe Lys Ser Val	Ser
85	90	95	
Ser Pro Glu Lys Pro	Lys Gly Gln Trp Leu	Asp Glu Ser Asp Ser	Asp
100	105	110	
Glu Glu Pro Ala Glu	Pro Ala Ser Ala Ala	Gly Ser Ala Ala Val	Ser
115	120	125	
Pro Val Lys Leu Leu	Pro Ser Gln Val Ala	Phe Ala Ser Gly Ser	Leu
130	135	140	
Leu Ser Pro Asp Pro	Thr Thr Ser Pro Met	Ser Pro Ser Ser Ser	Ser
145	150	155	160
Ala Ser Gly His Glu	Asp Pro Leu Gly Ile	Met Gly Val Asn Arg	Arg
165	170	175	
Arg Ser Leu Leu Glu	Gly Val Ser Leu Asp	Val Pro Ser His Ile	Asp
180	185	190	
Ser Lys Val Arg Ile	Ser Pro Val Pro Tyr	Leu Arg Leu Lys Lys	Ser
195	200	205	
Arg Ala Thr Lys Ala	Gln Trp Val Lys Glu	Lys Gly Arg Leu Pro	Phe

210 215 220

Leu Thr Val Tyr Arg Ala His Met Met Leu Met Thr Val Ile Cys Ile 230 235 Leu Ala Val Asp Phe Glu Val Phe Pro Arg Trp Gln Gly Lys Cys Glu 245 250 Asp Phe Gly Thr Ser Leu Met Asp Val Gly Val Gly Ser Phe Val Phe 260 Ser Leu Gly Leu Val Ser Thr Lys Ser Leu Ser Pro Pro Pro Thr 280 Pro Thr Pro Ser Ser Pro Ala Leu Asn Ser His Ile Ile Pro Leu Thr 295 Pro Ser Pro Phe Thr Ser Ile Leu Ile Ser Leu Arg Lys Ser Ile Pro 310 315 Ile Leu Val Leu Gly Phe Ile Arg Leu Ile Met Val Lys Gly Ser Asp Tyr Pro Glu His Val Thr Glu Tyr Gly Val His Trp Asn Phe Phe 345 Thr Leu Ala Leu Val Pro Val Leu Ala Val Gly Ile Arg Pro Leu Thr Gln Trp Leu Arg Trp Ser Val Leu Gly Val Ile Ile Ser Leu Leu His 375 Gln Leu Trp Leu Thr Tyr Tyr Leu Gln Ser Ile Val Phe Ser Phe Gly 395 Arg Ser Gly Ile Phe Leu Ala Asn Lys Glu Gly Phe Ser Ser Leu Pro 405 410 Gly Tyr Leu Ser Ile Phe Leu Ile Gly Leu Ser Ile Gly Asp His Val 420 425 Leu Arg Leu Ser Leu Pro Pro Arg Arg Glu Arg Val Val Ser Glu Thr Asn Glu Glu His Glu Gln Ser His Phe Glu Arg Lys Lys Leu Asp Leu Ile Met Glu Leu Ile Gly Tyr Ser Leu Gly Trp Trp Ala Leu Leu Gly 475 Gly Trp Ile Trp Ala Gly Gly Glu Val Ser Arg Arg Leu Ala Asn Ala Pro Tyr Val Phe Trp Val Ala Ala Tyr Asn Thr Thr Phe Leu Leu Gly 505 Tyr Leu Leu Leu Thr His Ile Ile Pro Ser Pro Thr Ser Ser Gln Thr 515 520 525 Ser Pro Ser Ile Leu Val Pro Pro Leu Leu Asp Ala Met Asn Lys Asn

Gly Leu Ala Ile Phe Leu Ala Ala Asn Leu Leu Thr Gly Leu Val Asn 545 550 Val Ser Met Lys Thr Met Tyr Ala Pro Ala Trp Leu Ser Met Gly Val 565 Leu Met Leu Tyr Thr Leu Thr Ile Ser Cys Val Gly Trp Ile Leu Lys 585 Gly Arg Arg Ile Lys Ile 595 <210> 15 <211> 35 <212> DNA <213> Artificial sequence <220> <223> Description of Artificial Sequence:an artificially synthesized primer sequence <400> 15 ggaattcatg tcgactttaa aacagagaaa agagg 35 <210> 16 <211> 34 <212> DNA <213> Artificial sequence <220> <223> Description of Artificial Sequence:an artificially

535

540

530

synthesized primer sequence

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